AMENDMENTS TO THE CLAIMS

1. (Currently Amended) A compound having the formula I

$$(R^3)_n = Q = N_1 + N_1 + N_2 + N_3 + N_4 + N_4 + N_5 + N_$$

(I)

wherein:

X1 is O or S:

X2 is a bond or C1-3alkylene;

P is C3.7cvcloalkvl or C4.7cvcloalkenvl;

R¹ is hydrogen, C₁₋₆alkyl, cyano, halogen and C₁₋₆alkylhalo, and one or more R¹ may be connected to each other or to one of the atoms that constitutes P to form a bridge or spirocyclo; R² is hydrogen, C₁₋₈alkyl, fluoromethyl, difluoromethyl, trifluoromethyl, methoxy,

fluoromethoxy, difluoromethoxy, trifluoromethoxy, $C_{0.3}$ alkylamino, $C_{0.3}$ alkylamino, $C_{0.3}$ alkylamino, $C_{0.3}$ alkylamino, $C_{0.3}$

C₁₋₃alkoxy, hydroxy or C₀₋₃alkyldimethylamino;

R⁴ is hydrogen, C₁₋₃alkyl, fluoromethyl, difluoromethyl, trifluoromethyl, methoxy,

fluoromethoxy, difluoromethoxy, trifluoromethoxy, C_{0-3} alkylamino C_{1-3} alkylamino,

€₀₋₃alkylhydroxy C₁₋₃alkoxy, hydroxy or C₀₋₃alkyldimethylamino;

Q is a saturated or partially unsaturated ring containing 4, 5, 6 or 7 atoms independently selected from C, S, O and N, and said ring may further contain groups independently selected from SO, SO, CO, evano and CS;

R3 is hydrogen, hydroxy, halogen, nitro, cyano, OC1,3alkylhalo, C1,3alkylhalo, C1,3alkyl,

 $C_{1\text{--}3}alkoxyC_{0\text{--}3}alkyl, C_{0\text{--}3}alkylOC_{2\text{--}4}alkanol, C_{1\text{--}3}alkanol, \underline{C_{0\text{--}3}alkylOhydroxyC_{2\text{--}4}alkyl,}$

 $\underline{hydroxyC_{1\text{--}3}alkyl}, amino, C_{1\text{--}3}alkylaminoC_{0\text{--}3}alkyl, (C_{1\text{--}3}alkyl)_2aminoC_{0\text{--}3}alkyl, amide,$

 C_{1-3} alkylamide C_{0-3} alkyl or $(C_{1-3}$ alkyl)₂amide C_{0-3} alkyl;

n is 0, 1, 2, 3 or 4; and

m is 0, 1, 2, 3 or 4:

2 ADM/mao

Application No. 10/766,948 Amendment dated January 22, 2007 Reply to Office Action of July 20, 2006

or N₁-oxides, or salts thereof, with the proviso that the following compound is excluded from formula 1: 3-amino 5,6,7,8 tetrahydrobenzof blpyrazine 2 (N-cyclohexy-t)carboxamide.

2. (Currently Amended) A compound having the formula I

$$(R^3)_n$$
 Q
 N_1
 N_1
 N_2
 N_3
 N_4
 $N_$

wherein:

X1 is O or S:

X2 is a bond or C1.3alkylene:

P is C3.7cycloalkyl or C4.7cycloalkenyl;

 R^1 is hydrogen, C_{1-6} alkyl, cyano, halogen and C_{1-6} alkylhalo, and one or more R^1 may be connected to each other or to one of the atoms that constitutes P to form a bridge or spirocyclo; R^2 is hydrogen, C_{1-3} alkyl, fluoromethyl, difluoromethyl, trifluoromethyl, methoxy,

fluoromethoxy, difluoromethoxy or trifluoromethoxy;

R4 is hydrogen;

Q is a <u>saturated or partially saturated</u> ring containing 4, 5, 6 or 7 atoms independently selected from C, S, O and N, which may be saturated or partially unsaturated and said ring may further contain groups independently selected from SO, SO₂, CO, cyano and CS;

 $R^{3} \ is \ hydrogen, \ hydroxy, \ halogen, \ nitro, \ OC_{1\text{--}3}alkylhalo, \ C_{1\text{--}3}alkylhalo, \ C_{1\text{--}3}alkylhal$

C₁₋₃alkoxyC₀₋₃alkyl, C₁₋₃alkanol, hydroxyC₁₋₃alkyl, cyano, amino or amide;

n is 0, 1, 2, 3 or 4; and

m is 0, 1, 2, 3 or 4;

or N₁-oxides, or salts thereof.

3 ADM/mao

Application No. 10/766,948 Amendment dated January 22, 2007

Reply to Office Action of July 20, 2006

3. (Original) The compound according to any one of claims 1 or 2, wherein P is C₃₋₇cycloalkyl

substituted with one or more R1, wherein R1 is hydrogen, C1-6alkyl, cyano, halogen or

C₁₋₆alkylhalo, and one or more R¹ may be connected to each other or to one of the atoms that

constitutes P to form a bridge or spirocyclo.

4. (Original) The compound according to claim 3, wherein P is C₅₋₇cycloalkyl substituted with

one or more R1, wherein R1 is methyl.

5. (Previously Presented) The compound according to any one of claims 1 or 2, wherein X¹ is

oxygen.

6. (Previously Presented) The compound according to any one of claims 1 or 2, wherein X² is a

bond.

7. (Previously Presented) The compound according to any one of claims 1 or 2, wherein R² is

hydrogen.

8. (Previously Presented) The compound according to any one of claims 1 or 2, wherein R⁴ is

hydrogen or methyl.

9. (Currently Amended) The compound according to any one of claims 1 or 2, wherein Q is a

saturated or partially unsaturated ring containing 5, 6 or 7 atoms independently selected from C,

O and N. which may be saturated or partially unsaturated.

10. (Previously Presented) The compound according to any one of claims 1 or 2, wherein R3 is

 $hydrogen,\,hydroxy,\,halogen,\,cyano,\,C_{1\text{--}3}alkyl\,or\,\,C_{1\text{--}3}alkoxyC_{0\text{--}3}alkyl.$

4 ADM/mao

11. (Previously Presented) The compound according to any one of claims 1 or 2 having a transrelationship between R^1 and X^2 on ring P, wherein P is cyclohexane, and R^1 and X^2 are attached to P at positions 4 and 1, respectively.

12. (Previously Presented) The compounds N-(trans-4-methylcyclohexyl)-5,6,7,8-tetrahydroquinoxaline-2-carboxamide,

N-(4,4-dimethylcyclohexyl)-5,6,7,8-tetrahydroquinoxaline-2-carboxamide, or salts thereof.

13. (Previously Presented) The compounds

N-(4,4-dimethylcyclohexyl)-3-methyl-5,6,7,8-tetrahydroquinoxaline-2-carboxamide,
8-methyl-N-(trans-4-methylcyclohexyl)-5,6,7,8-tetrahydroquinoxaline-2-carboxamide,
7-hydroxy-5,7-dimethyl-N-(trans-4-methylcyclohexyl)-6,7-dihydro-5H-cyclopenta[b]pyrazine-2-carboxamide

N-(trans-4-methylcyclohexyl)-6,7,8,9-tetrahydro-5H-cyclohepta[b]pyrazine-2-carboxamide,
7-methyl-N-(trans-4-methylcyclohexyl)-5,6,7,8-tetrahydroquinoxaline-2-carboxamide,
6-methyl-N-(trans-4-methylcyclohexyl)-5,6,7,8-tetrahydroquinoxaline-2-carboxamide,
N-(trans-4-methylcyclohexyl)-6,7-dihydro-5H-cyclopenta[b]pyrazine-2-carboxamide,
N-(trans-4-methylcyclohexyl)-7,8-dihydro-5H-pyrano[3,4-b]pyrazine-2-carboxamide,
N-(trans-4-methylcyclohexyl)-7,8-dihydro-5H-pyrano[3,4-b]pyrazine-3-carboxamide,
7-hydroxy-N-(trans-4-methylcyclohexyl)-5,6,7,8-tetrahydroquinoxaline-2-carboxamide,
6-hydroxy-N-(trans-4-methylcyclohexyl)-5,6,7,8-tetrahydroquinoxaline-2-carboxamide,
N-(4,4-dimethylcyclohexyl)-5,6,7,8-tetrahydroquinoxaline-2-carboxamide 4-oxide and
6,7-dimethyl-N-(4-methylcyclohexyl)-6,7-dihydro-5H-cyclopenta[b]pyrazine-2-carboxamide,
or salts thereof.

14. (Withdrawn and Previously Presented) A pharmaceutical composition comprising as active ingredient a therapeutically effective amount of the compound according to any one of claims 1

or 2, in association with one or more pharmaceutically acceptable diluent, excipients and/or inert carrier.

15. (Withdrawn) The pharmaceutical composition according to claim 14, for use in the treatment of Group I mGluR mediated disorders.

16.-18. (Cancelled)

19. (Withdrawn and Previously Presented) A method of treatment of Group I mGluR mediated disorders, comprising administering to a mammal, including man in need of such treatment, a therapeutically effective amount of the compound according to any one of claims 1 or 2.

20. (Withdrawn) The method according to claim 19, for use in treatment of neurological disorders.

21. (Withdrawn) The method according to claim 19, for use in treatment of psychiatric disorders.

22. (Withdrawn) The method according to claim 19, for use in treatment of chronic and acute pain disorders.

23. (Withdrawn) The method according to claim 19, for use in treatment of gastrointestinal disorders.

24. (Withdrawn) A method for inhibiting activation of Group I mGluR receptors, comprising treating a cell containing said receptor with an effective amount of the compound according to claim 1 or 2

Amendment dated January 22, 2007 Reply to Office Action of July 20, 2006

25. (Withdrawn) Processes for the preparation of the compound according to claim 1 or 2, wherein $P, Q, X^1, X^2, R^1, R^2, R^3, R^4$, m and n are, unless otherwise specified, defined as in formula I, comprising of:

A
$$(R^{3})_{n} \xrightarrow{X^{1}} O \xrightarrow{R^{y}} HN \xrightarrow{X^{2}} P \xrightarrow{(R^{1})_{m}} (VIII)$$

$$(R^{3})_{n} \xrightarrow{Q} N \xrightarrow{X^{1}} X^{2} \xrightarrow{Q} R^{1}$$

$$(R^{3})_{n} \xrightarrow{Q} N \xrightarrow{X^{1}} X^{2} \xrightarrow{Q} R^{1}$$

$$(I)$$

reacting a compound of formula VII, wherein R^y is H, with an activating agent followed by the treatment of the resulting acid halide, or otherwise to nucleophiles activated acid derivative, with an amine of formula XIV, to obtain the compound of formula I, alternatively.

7 ADM/mao

Reply to Office Action of July 20, 2006

В

reacting an amine of formula XIV with the compound of formula VII, wherein R^y is H, to obtain the compound of formula I, or

C

$$(R^3)_n \qquad X^1 \qquad X^1 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^1 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^1 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad Y^1 \qquad Y^2 \qquad Y^1 \qquad$$

reacting a compound of formula VIa or the N_I -oxide thereof, wherein R^x is $C_{I-\alpha}$ alkyl, with the appropriate amine such as the compound of formula XIV, to obtain the compound of formula I,

or,

D

direct condensation of intermediates of formula IV and XVb, to obtain the compound of formula I.

- 26. (Withdrawn) Compounds
- 5.6.7.8-tetrahydro-quinoxaline-2-carboxylic acid methyl ester and
- 5,6,7,8-tetrahydro-quinoxaline-2-carboxylic acid.
- 27. (Withdrawn) Compounds
- 3-methyl-5,6,7,8-tetrahydro-quinoxaline-2-carboxylic acid ethyl ester,
- 3-methyl-5,6,7,8-tetrahydro-quinoxaline-2-carboxylic acid,
- 2,3-diamino-N-(4-methyl-cyclohexyl)-propionamide,
- 4-(tert-butyl-diphenyl-silanyloxy)-cyclohexane-1,2-dione,
- 6,7-dimethyl-6,7-dihydro-5H-cyclopentapyrazine-2-carboxylic acid methyl ester,
- 5,6,7,8-tetrahydro-quinoxaline-2-carboxylic acid methyl ester and
- 5.6,7,8-tetrahydro-quinoxaline-2-carboxylic acid.

9 ADM/mao

plication No. 10/766,948 Docket No.: 5999-0524PUS3

28. (Withdrawn) The compounds according to claims 26 and 27, for use as an intermediate in the preparation of the compound according to claim 1.

- 29. (Previously Presented) The compound according to claim 3, wherein X¹ is oxygen.
- 30. (Previously Presented) The compound according to claim 4, wherein X1 is oxygen.
- 31. (Previously Presented) The compound according to claim 3, wherein X² is a bond.
- (Previously Presented) The compound according to claim 4, wherein X² is a bond.
- (Previously Presented) The compound according to claim 5, wherein X² is a bond.
- 34. (Previously Presented) The compound according to claim 3, wherein R² is hydrogen.
- 35. (Previously Presented) The compound according to claim 4, wherein R² is hydrogen.
- 36. (Previously Presented) The compound according to claim 5, wherein R² is hydrogen.
- 37. (Previously Presented) The compound according to claim 6, wherein R² is hydrogen.
- 38. (Previously Presented) The compound according to claim 3, wherein R⁴ is hydrogen or methyl.
- 39. (Previously Presented) The compound according to claim 4, wherein \mathbb{R}^4 is hydrogen or methyl.
- 40. (Previously Presented) The compound according to claim 5, wherein \mathbb{R}^4 is hydrogen or methyl.

41. (Previously Presented) The compound according to claim 6, wherein R⁴ is hydrogen or methyl.

42. (Previously Presented) The compound according to claim 7, wherein R⁴ is hydrogen or methyl.

43. (Currently Amended) The compound according to claim 3, wherein Q is a <u>saturated or partially unsaturated</u> ring containing 5, 6 or 7 atoms independently selected from C, O and N₂ which may be saturated or partially unsaturated.

44. (Currently Amended) The compound according to claim 4, wherein Q is a <u>saturated or partially unsaturated</u> ring containing 5, 6 or 7 atoms independently selected from C, O and Nawhich may be <u>saturated or partially unsaturated</u>.

45. (Currently Amended) The compound according to claim 5, wherein Q is a <u>saturated or partially unsaturated</u> ring containing 5, 6 or 7 atoms independently selected from C, O and N₂ which may be saturated or partially unsaturated.

46. (Currently Amended) The compound according to claim 6, wherein Q is a <u>saturated or partially unsaturated</u> ring containing 5, 6 or 7 atoms independently selected from C, O and N_c which may be saturated or partially unsaturated.

47. (Currently Amended) The compound according to claim 7, wherein Q is a saturated or partially unsaturated ring containing 5, 6 or 7 atoms independently selected from C, O and $N_{\rm s}$ which may be saturated or partially unsaturated.

- 48. (Currently Amended) The compound according to claim 8, wherein Q is a saturated or partially unsaturated ring containing 5, 6 or 7 atoms independently selected from C, O and N_τ which may be saturated or partially unsaturated.
- (Previously Presented) The compound according to claim 3, wherein R³ is hydrogen, hydroxy, halogen, cyano, C₁₋₃alkyl or C₁₋₃alkoxyC₀₋₃alkyl.
- 50. (Previously Presented) The compound according to claim 4, wherein R^3 is hydrogen, hydroxy, halogen, cyano, $C_{1.3}$ alkyl or $C_{1.3}$ alkoxy $C_{0.3}$ alkyl.
- 51. (Previously Presented) The compound according to claim 5, wherein R³ is hydrogen, hydroxy, halogen, cyano, C_{1.3}alkyl or C_{1.3}alkyyC_{0.3}alkyl.
- (Previously Presented) The compound according to claim 6, wherein R³ is hydrogen, hydroxy, halogen, cyano, C_{1,3}alkyl or C_{1,3}alkoyC_{0,3}alkyl.
- 53. (Previously Presented) The compound according to claim 7, wherein R³ is hydrogen, hydroxy, halogen, cyano, C_{1.3}alkyl or C_{1.3}alkoxyC_{0.3}alkyl.
- 54. (Previously Presented) The compound according to claim 8, wherein R³ is hydrogen, hydroxy, halogen, evano, C_{1.3}alkyl or C_{1.3}alkoxyC_{0.3}alkyl.
- 55. (Previously Presented) The compound according to claim 9, wherein \mathbb{R}^3 is hydrogen, hydroxy, halogen, cyano, $C_{1,3}$ alkyl or $C_{1,3}$ alkyl or $C_{1,3}$ alkyl.
- 56. (Previously Presented) The compound according to claim 3 having a trans-relationship between R¹ and X² on ring P, wherein P is cyclohexane, and R¹ and X² are attached to P at positions 4 and 1, respectively.

57. (Previously Presented) The compound according to claim 4 having a trans-relationship between R¹ and X² on ring P, wherein P is cyclohexane, and R¹ and X² are attached to P at

positions 4 and 1, respectively.

58. (Previously Presented) The compound according to claim 5 having a trans-relationship between R^1 and X^2 on ring P, wherein P is cyclohexane, and R^1 and X^2 are attached to P at

positions 4 and 1, respectively.

59. (Previously Presented) The compound according to claim 6 having a trans-relationship

between R^1 and X^2 on ring P, wherein P is cyclohexane, and R^1 and X^2 are attached to P at

positions 4 and 1, respectively.

60. (Previously Presented) The compound according to claim 7 having a trans-relationship

between R1 and X2 on ring P, wherein P is cyclohexane, and R1 and X2 are attached to P at

positions 4 and 1, respectively.

61. (Previously Presented) The compound according to claim 8 having a trans-relationship

between R1 and X2 on ring P, wherein P is cyclohexane, and R1 and X2 are attached to P at

positions 4 and 1, respectively.

62. (Previously Presented) The compound according to claim 9 having a trans-relationship

between R1 and X2 on ring P, wherein P is cyclohexane, and R1 and X2 are attached to P at

positions 4 and 1, respectively.

63. (Previously Presented) The compound according to claim 10 having a trans-relationship

between R^1 and X^2 on ring P, wherein P is cyclohexane, and R^1 and X^2 are attached to P at

positions 4 and 1, respectively.

Application No. 10/766,948 Amendment dated January 22, 2007 Reply to Office Action of July 20, 2006

64. (New) The compound according to Claim I, wherein R^4 is hydrogen, $C_{1.3}$ alkyl, fluoromethyl, difluoromethyl, trifluoromethyl, methoxy, fluoromethoxy, difluoromethoxy, trifluoromethoxy, $C_{1.3}$ alkylamino, $C_{1.3}$ alkoxy, hydroxy.

65. (New) The compound according to any one of Claims 1 or 2, wherein Q is cyclohexpl, cyclohexenyl, cyclopentryl, cyclopentenyl, imidazolidinyl, imidazolinyl, morpholinyl, piperazinyl, piperidyl, piperidonyl, pyrazolidinyl, pyrazolinyl, pyrrolinyl, tetrahydropyranyl or thiomorpholinyl.

14 ADM/mao